WO 03/064421

(I)

Claims

1. A compound of formula (I) or a pharmaceutically acceptable derivative thereof:

$$R^{1} \xrightarrow{Z^{2}} Z^{3} \xrightarrow{N} Z^{4}$$

$$R^{2} \xrightarrow{Z^{3}} R^{3}$$

$$R^{3} \xrightarrow{R^{3}} R^{3}$$

wherein:

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one of Z^1 , Z^2 , Z^3 , Z^4 and Z^5 is N, one is CR^{1a} and the remainder are CH, or one of Z^1 , Z^2 , Z^3 , Z^4 and Z^5 is CR^{1a} and the remainder are CH;

 R^1 and R^{1a} are independently hydrogen; hydroxy; (C_{1-6}) alkoxy optionally substituted by (C_{1-6}) alkoxy, amino, piperidyl, guanidino or amidino any of which is optionally N-substituted by one or two (C_{1-6}) alkyl, acyl or (C_{1-6}) alkylsulphonyl groups, CONH₂, hydroxy, (C_{1-6}) alkylthio, heterocyclylthio, heterocyclyloxy, arylthio, aryloxy, acylthio, acyloxy or (C_{1-6}) alkylsulphonyloxy; (C_{1-6}) alkoxy-substituted (C_{1-6}) alkyl; halogen; (C_{1-6}) alkyl; (C_{1-6}) alkylthio; trifluoromethyl; trifluoromethoxy; nitro; azido; acyl; acyloxy; acylthio; (C_{1-6}) alkylsulphonyl; (C_{1-6}) alkylsulphoxide; arylsulphonyl; arylsulphoxide or an amino, piperidyl, guanidino or amidino group optionally N-substituted by one or two (C_{1-6}) alkyl, acyl or (C_{1-6}) alkylsulphonyl groups;

or when Z⁵ is CR^{1a}, R^{1a} may instead be cyano, hydroxymethyl or carboxy;

or R^1 and R^{1a} on adjacent positions may together form ethylenedioxy;

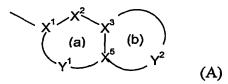
provided that when none of Z^1 , Z^2 , Z^3 , Z^4 and Z^5 is N, then R^1 is not hydrogen;

 R^2 is hydrogen, or (C_{1-4}) alkyl or (C_{2-4}) alkenyl optionally substituted with 1 to 3 groups selected from:

amino optionally substituted by one or two (C₁₋₄)alkyl groups; carboxy; (C₁₋₄)alkoxycarbonyl; (C₁₋₄)alkylcarbonyl; (C₂₋₄)alkenyloxycarbonyl; (C₂₋₄)alkenylcarbonyl; aminocarbonyl wherein the amino group is optionally substituted by hydroxy, (C₁₋₄)alkyl, hydroxy(C₁₋₄)alkyl, aminocarbonyl(C₁₋₄)alkyl, (C₂₋₄)alkenyl,

 $\label{eq:continuous} $$(C_{1-4})$ alkylsulphonyl, trifluoromethylsulphonyl, (C_{2-4})$ alkenylsulphonyl, (C_{1-4})$ alkylcarbonyl, (C_{2-4})$ alkenyloxycarbonyl or (C_{2-4})$ alkenylcarbonyl; cyano; tetrazolyl; 2-oxo-oxazolidinyl optionally substituted by R^{10}; 3-hydroxy-3-cyclobutene-1,2-dione-4-yl; 2,4-thiazolidinedione-5-yl; tetrazol-5-$

- ylaminocarbonyl; 1,2,4-triazol-5-yl optionally substituted by R¹⁰; 5-oxo-1,2,4-oxadiazol-3-yl; halogen; (C₁₋₄)alkylthio; trifluoromethyl; hydroxy optionally substituted by (C₁₋₄)alkyl, (C₂₋₄)alkenyl, (C₁₋₄)alkoxycarbonyl, (C₁₋₄)alkylcarbonyl, (C₂₋₄)alkenyloxycarbonyl, (C₂₋₄)alkenylcarbonyl; oxo; (C₁₋₄)alkylsulphonyl; (C₂₋₄)alkenylsulphonyl; or (C₁₋₄)aminosulphonyl wherein the amino group is optionally substituted by (C₁₋₄)alkyl or (C₂₋₄)alkenyl;
 - R^3 is in the 2-, 3- or 4-position and is trifluoromethyl or is in the 2-position and is oxo; or R^3 is in the 3-position and is fluorine or amino wherein the amino group is optionally substituted by: hydroxy; (C_{1-6}) alkylsulphonyl; trifluoromethylsulphonyl; (C_{2-})
- 6)alkenylsulphonyl; (C_{1-6}) alkylcarbonyl; (C_{2-6}) alkenylcarbonyl; (C_{1-6}) alkoxycarbonyl; (C_{2-6}) alkenyloxycarbonyl; (C_{1-6}) alkyl; or (C_{2-6}) alkenyl; wherein a (C_{1-6}) alkyl or (C_{2-6}) alkenyl moiety may be optionally substituted with up to 2 groups R^{12} independently selected from:
- halogen; (C₁₋₆)alkylthio; trifluoromethyl; cyano; carboxy; tetrazolyl; 2-oxo-oxazolidinyl; 3-hydroxy-3-cyclobutene-1,2-dione-4-yl; 2,4-thiazolidinedione-5-yl; tetrazol-5-ylaminocarbonyl; 1,2,4-triazol-5-yl optionally substituted by R¹⁰; or 5-oxo-1,2,4-oxadiazol-3-yl; (C₁₋₆)alkoxycarbonyl; (C₁₋₆)alkylcarbonyl; (C₂₋₆)alkenylcarbonyl; hydroxy optionally substituted by (C₁₋₆)alkyl, (C₂₋₆)alkenyl, (C₁₋₆)alkylcarbonyl, (C₂₋₆)alkenylcarbonyl or aminocarbonyl
- wherein the amino group is optionally substituted by (C₁₋₆)alkyl, (C₂₋₆)alkenyl; amino optionally mono- or disubstituted by (C₁₋₆)alkoxycarbonyl, (C₁₋₆)alkylcarbonyl, (C₂₋₆)alkenylcarbonyl, (C₁₋₆)alkyl, (C₂₋₆)alkenyl, (C₁₋₆)alkylsulphonyl, (C₂₋₆)alkenylsulphonyl or aminocarbonyl wherein the amino group is optionally substituted by (C₁₋₆)alkyl or (C₂₋₆)alkenyl;
- in addition when R³ is disubstituted with a hydroxy or amino containing substituent and carboxy containing substituent these may together form a cyclic ester or amide linkage, respectively;
 - R⁴ is a group -U-R⁵ where
- U is selected from CO, SO₂ and CH₂ and
 R⁵ is an optionally substituted bicyclic carbocyclic or heterocyclic ring system (A):



containing up to four heteroatoms in each ring in which

ring (a) is aromatic and ring (b) is non-aromatic;

 X^1 is C or N;

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 X^2 is N, NR¹³, O, S(O)_x, CO or CR¹⁴;

 X^3 and X^5 are independently N or C;

 Y^1 is a 0 to 4 atom linker group each atom of which is independently selected from N, NR¹³, O, S(O)_x, CO and CR¹⁴;

Y² is a 2 to 6 atom linker group, each atom of Y² being independently selected from N, NR¹³, O, S(O)_x, CO, CR¹⁴ and CR¹⁴R¹⁵; each of R¹⁴ and R¹⁵ is independently selected from: H; (C₁₋₄)alkylthio; halo; carboxy(C₁₋₄)alkyl; halo(C₁₋₄)alkoxy; halo(C₁₋₄)alkyl; (C₁₋₄)alkyl; (C₂₋₄)alkenyl; (C₁₋₄)alkoxycarbonyl; formyl; (C₁₋₄)alkylcarbonyl; (C₂₋₄)alkenyloxycarbonyl; (C₂₋₄)alkenylcarbonyl; (C₁₋₄)alkylcarbonyloxy; (C₁₋₄)alkoxycarbonyl(C₁₋₄)alkyl; hydroxy; hydroxy(C₁₋₄)alkyl; mercapto(C₁₋₄)alkyl; (C₁₋₄)alkoxy; trifluoromethoxy; nitro; cyano; carboxy; amino or aminocarbonyl optionally substituted as for corresponding substituents in R³; (C₁₋₄)alkylsulphonyl; (C₂₋₄)alkenylsulphonyl; or aminosulphonyl wherein the amino group is optionally mono- or di-substituted by (C₁₋₄)alkyl or (C₂₋₄)alkenyl; aryl; aryl(C₁₋₄)alkyl; aryl(C₁₋₄)alkoxy;

each R^{13} is independently H; trifluoromethyl; (C_{1-4}) alkyl optionally substituted by hydroxy, (C_{1-6}) alkoxy, (C_{1-6}) alkylthio, halo or trifluoromethyl; (C_{2-4}) alkenyl; aryl; aryl (C_{1-4}) alkyl; arylcarbonyl; heteroarylcarbonyl; (C_{1-4}) alkoxycarbonyl; (C_{1-4}) alkylcarbonyl; formyl; (C_{1-6}) alkylsulphonyl; or aminocarbonyl wherein the amino group is optionally substituted by (C_{1-4}) alkoxycarbonyl, (C_{1-4}) alkylcarbonyl, (C_{2-4}) alkenyloxycarbonyl, (C_{2-4}) alkenylcarbonyl, (C_{2-4}) alkenylcarbonyl;

each x is independently 0, 1 or 2

n is 0 and AB is NR¹¹CO, CO-CR⁸R⁹, CR⁶R⁷-CO, NHR¹¹SO₂, CR⁶R⁷-SO₂ or CR⁶R⁷-CR⁸R⁹, provided that R⁸ and R⁹ are not optionally substituted hydroxy or amino and R⁶ and R⁸ do not represent a bond: or n is 1 and AB is NR¹¹CO, CO-CR⁸R⁹, CR⁶R⁷-CO, NR¹¹SO₂, CONR¹¹, CR⁶R⁷-CR⁸R⁹, O-CR⁸R⁹ or NR¹¹-CR⁸R⁹;

each of R⁶, R⁷, R⁸ and R⁹ is independently selected from: hydrogen; (C₁₋₆)alkoxy; (C₁₋₆)alkylthio; halo; trifluoromethyl; azido; (C₁₋₆)alkyl; (C₂₋₆)alkenyl; (C₁₋₆)alkoxycarbonyl; (C₁₋₆)alkylcarbonyl; (C₂₋₆)alkenyloxycarbonyl; (C₂₋₆)alkenylcarbonyl; hydroxy, amino or aminocarbonyl optionally substituted as for corresponding substituents in R³; (C₁₋₆)alkylsulphonyl; (C₂₋₆)alkenylsulphonyl; or aminosulphonyl wherein the amino group is optionally substituted by (C₁₋₆)alkyl or (C₂₋₆)alkenyl; or when n=1 R⁶ and R⁸ together represent a bond and R⁷ and R⁹ are as above defined; or R⁶ and R⁷ or R⁸ and R⁹ together represent oxo;

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 $\rm R^{10}$ is selected from (C1-4)alkyl; (C2-4)alkenyl and aryl any of which may be optionally substituted by a group $\rm R^{12}$ as defined above; carboxy; aminocarbonyl wherein the amino group is optionally substituted by hydroxy, (C1-6)alkyl, (C2-6)alkenyl, (C1-6)alkylsulphonyl, trifluoromethylsulphonyl, (C2-6)alkenylsulphonyl, (C1-6)alkoxycarbonyl, (C1-6)alkylcarbonyl, (C2-6)alkenyloxycarbonyl or (C2-6)alkenyloxycarbonyloxycarbonyl or (C2-6)alkenyloxycarbonyl or (C2-6)alkenyloxycarbonyl or (C2-6)alkenyloxycarbonyl or (C2-6)a

6)alkoxycarbonyl, (C₁₋₆)alkylcarbonyl, (C₂₋₆)alkenyloxycarbonyl or (C₂₋₆)alkenylcarbonyl and optionally further substituted by (C₁₋₆)alkyl or (C₂₋₆)alkenyl; and

 R^{11} is hydrogen; trifluoromethyl, (C_{1-6}) alkyl; (C_{2-6}) alkenyl; (C_{1-6}) alkoxycarbonyl; (C_{1-6}) alkylcarbonyl; or aminocarbonyl wherein the amino group is optionally substituted by (C_{1-6}) alkoxycarbonyl, (C_{1-6}) alkylcarbonyl, (C_{2-6}) alkenyloxycarbonyl, (C_{2-6}) alkenylcarbonyl, (C_{1-6}) alkyl or (C_{2-6}) alkenyl and optionally further substituted by (C_{1-6}) alkyl or (C_{2-6}) alkenyl;

or where one of R³ and R⁶, R⁷, R⁸ or R⁹ contains a carboxy group and the other contains a hydroxy or amino group they may together form a cyclic ester or amide linkage.

2. A compound according to claim 1 wherein Z^5 is CH, C-Cl or N, Z^3 is CH or CF and Z^1 , Z^2 and Z^4 are each CH, or Z^1 is N, Z^3 is CH and Z^2 and Z^4 are each CH and Z^5 is CH or C-Cl.

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- 3. A compound according to any preceding claim wherein R^1 is methoxy and R^{1a} is H or when Z^3 is CR^{1a} it may be C-F or when Z^5 is CR^{1a} it may be C-F or C-Cl.
- 4. A compound according to any preceding claim wherein R² is hydrogen, carboxymethyl, hydroxyethyl, aminocarbonylmethyl, ethoxycarbonylmethyl, ethoxycarbonylallyl or carboxyallyl.

5. A compound according to any preceding claim wherein \mathbb{R}^3 is \mathbb{CF}_3 , fluoro, oxo or amino unsubstituted or substituted by (\mathbb{C}_{1-6}) alkyl or (\mathbb{C}_{2-6}) alkenyl.

- 6. A compound according to any preceding claim wherein n is 0 and either A is CH₂ or CHOH and B is CH₂ or A is NH and B is CO.
 - 7. A compound according to any preceding claim wherein -U- is -CH₂-.
- 8. A compound according to any preceding claim wherein in the heterocyclic ring (A) ring (a) is selected from optionally substituted benzo and pyrido and Y² has 3-5 atoms including a heteroatom bonded to X⁵ selected from NR¹³, O or S, where R¹³ is other than hydrogen, and NHCO bonded via N to X³, or O or NH bonded to X³.
- 9. A compound according to any one of claims 1 to 6 wherein R⁵ is selected from:
 4H-benzo[1,4] oxazin-3-one-6-yl
 4H-benzo[1,4] thiazin-3-one-6-yl

2,3-dihydro-[1,4]dioxino[2,3-c]pyridin-7-yl 3-oxo-3,4-dihydro-2H-pyrido[3,2-b][1,4]thiazin-6-yl

3-oxo-3,4-dihydro-2H-pyrido[3,2-b][1,4]oxazin-6-yl

7-chloro-3-oxo-3,4-dihydro-2H-pyrido[3,2-b][1,4]oxazin-6-yl

7-chloro-3-oxo-3,4-dihydro-2H-pyrido[3,2-b][1,4]thiazin-6-yl.

- 10. A compound according to claim 1 selected from:
- $6-(\{2S,4S\}-1-[(R)-2-Hydroxy-2-(6-methoxy-[1,5]naphthyridin-4-yl)ethyl]-2-$
- 25 (trifluoromethyl)piperidin-4-ylamino}methyl)-4*H*-benzo[1,4]thiazin-3-one; 6-({(3*R*,4*S*)-1-[(*R*)-2-Hydroxy-2-(6-methoxy-[1,5]naphthyridin-4-yl)ethyl]-3-(trifluoromethyl)piperidin-4-ylamino}methyl)-4*H*-benzo[1,4]thiazin-3-one; 6-({1-[(*R*)-2-Hydroxy-2-(6-methoxy-[1,5]naphthyridin-4-yl)ethyl]-4-

(trifluoromethyl)piperidin-4-ylamino}methyl)-4H-benzo[1,4]thiazin-3-one;

6-({1-[(R)-2-Hydroxy-2-(6-methoxyquinolin-4-yl)ethyl]-2-oxopiperidin-4-ylamino}methyl)-4H-benzo[1,4]thiazin-3-one;
6-[({(3S,4R)-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-

ylamino}methyl)-4H-benzo[1,4]thiazin-3-one and 6-[({(3R,4S)-3-fluoro-1-[(R)-2-

hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-

35 benzo[1,4]thiazin-3-one; 6-({cis-3-Fluoro-1-[(R)-2-hydroxy-2-(

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6-({cis-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one Diastereoisomer 1;

6-({cis-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]-piperidin-4-ylamino}-methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one Diastereoisomer 2; 7-Chloro-6-({cis 3-fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one Diastereoisomer 1;

- 7-Chloro-6-({cis-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one Diastereoisomer 2; 6-({cis-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]oxazin-3-one Diastereoisomer 1; 6-({cis-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-
- ylamino}methyl)-4H-pyrido[3,2-b][1,4]oxazin-3-one Diastereoisomer 2; 7-Chloro-6-[($\{(3S,4R)$ -3-fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one and 7-chloro-6-[($\{(3R,4S)$ -3-fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one;
- 7-Fluoro-6-({(3S,4R)-3-fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one and 7-fluoro-6-[({(3R,4S)-3-fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one;
 - $7-(\{(3S,4R)-3-\text{fluoro}-1-[(R)-2-\text{hydroxy}-2-(6-\text{methoxyquinolin}-4-\text{yl})-\text{ethyl}]$ piperidin-4-
- ylamino}methyl)-1H-pyrido[2,3-b][1,4]thiazin-2-one and 7-($\{(3R,4S)$ -3-fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-1H-pyrido[2,3-b][1,4]thiazin-2-one;
 - 7-Chloro-6- $[({(3S,4R)-3-fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]oxazin-3-one and 7-chloro-6-$
- [({(3R,4S)-3-fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]oxazin-3-one;
 6-[({(3S,4S)-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one and 6-[({(3R,4R)-3-fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one and 6-[(4,4R)-3-fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one and 6-[(4,4R)-3-fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-ylamino)]methyl]
- b][1,4]thiazin-3-one;
 6-[({(3S,4S)-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]oxazin-3-one and 6-[({(3R,4R)-3-fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]oxazin-3-one;
- 7-Fluoro-6-[({(3S,4S)-3-fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one and 7-Fluoro-6-[({(3R,4R)-3-fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-

ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one; 6-[(${(3S,4S)-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-<math>1H$ -pyrido[2,3-b][1,4]thiazin-3-one and 6-[(${(3R,4R)-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-<math>1H$ -

- pyrido[2,3-b][1,4]thiazin-3-one; 6-({cis-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxy-[1,5]naphthyridin)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]oxazin-3-one Diastereoisomer 1; 6-({cis-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxy-[1,5]naphthyridin)-ethyl]piperidin-4-
 - 6-({cis-3-Fluoro-1-[(*R*)-2-hydroxy-2-(6-methoxy-[1,5]naphthyridin)-ethyl]piperidin-4-ylamino}methyl)-4*H*-pyrido[3,2-*b*][1,4]oxazin-3-one Diastereoisomer 2;
- 6-({cis-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxy-[1,5]naphthyridin)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one Diastereoisomer 1;
 6-({cis-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxy-[1,5]naphthyridin)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one Diastereoisomer 2;
 7-Chloro-6-({cis-3-fluoro-1-[(R)-2-hydroxy-2-(6-methoxy-[1,5]naphthyridin)-
- ethyl]piperidin-4-ylamino}methyl)-4*H*-pyrido[3,2-*b*][1,4]oxazin-3-one Diastereoisomer 1;
 7-Chloro-6-({cis-3-fluoro-1-[(*R*)-2-hydroxy-2-(6-methoxy-[1,5]naphthyridin)-ethyl]piperidin-4-ylamino}methyl)-4*H*-pyrido[3,2-*b*][1,4]oxazin-3-one Diastereoisomer 2;
- 6-({cis-3-Fluoro-1-[(R)-2-hydroxy-2-(8-fluoro-6-methoxy-quinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]oxazin-3-one Diastereoisomer 1; 6-({cis-3-Fluoro-1-[(R)-2-hydroxy-2-(8-fluoro-6-methoxy-quinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one Diastereoisomer 1; 6-({(3R,4S)-1-[2-(3-Chloro-6-methoxy-quinolin-4-yl)-ethyl]-3-fluoro-piperidin-4-
- ylamino}methyl)-4*H*-pyrido[3,2-*b*][1,4]thiazin-3-one and 6-({(3S,4R)-1-[2-(3-Chloro-6-methoxy-quinolin-4-yl)-ethyl]-3-fluoro-piperidin-4-ylamino}methyl)-4*H*-pyrido[3,2-*b*][1,4]thiazin-3-one;
 - $6-(\{(3R,4S)-3-Fluoro-1-[2-(6-methoxy-[1,5]naphthyridin-4-yl)-ethyl]-piperidin-4-ylamino\}-methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one and 6-(\{(3S,4R)-3-fluoro-1-[2-(6-methoxy-[1,5]naphthyridin-4-yl)-ethyl]-piperidin-4-ylamino\}-methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one and 6-(\{(3S,4R)-3-fluoro-1-[2-(6-methoxy-[1,5]naphthyridin-4-yl)-ethyl]-piperidin-4-ylamino\}-methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one and 6-(\{(3S,4R)-3-fluoro-1-[2-(6-methoxy-[1,5]naphthyridin-4-yl)-ethyl]-piperidin-4-ylamino}- 0-1-[2-(6-methoxy-[1,5]naphthyridin-4-yl)-ethyl]-piperidin-4-ylamino}- 0-1-[2-(6-methoxy-[1,5]naphthyridin-4-yl)-ethyl]-piperidin-4-ylamino}- 0-1-[2-(6-methoxy-[1,5]naphthyridin-4-yl)-ethyl]-piperidin-4-ylamino}- 0-1-[2-(6-methoxy-[1,5]naphthyridin-4-yl)-ethyl]-piperidin-4-ylamino}- 0-1-[2-(6-methoxy-[1,5]naphthyridin-4-yl)-ethyl]-piperidin-4-ylamino}- 0-1-[2-(6-methoxy-[1,5]naphthyridin-4-yl)-ethyl]-piperidin-4-ylamino}- 0-1-[2-(6-methoxy-[1,5]naphthyridin-4-ylamino}- 0-1-[2-(6-metho$
- methoxy-[1,5]naphthyridin-4-yl)-ethyl]-piperidin-4-ylamino}-methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one;
 6-[({(3S,4R)-3-Fluoro-1-[(S)-2-hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one and 6-[({(3R,4S)-3-fluoro-1-[(S)-2-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one and 6-[({(3R,4S)-3-[(S)-2-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one and 6-[({(3R,4S)-3-[(S)-2-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-[(S)-2-ylamino]methyl]-4H-pyrido[3,2-b][1,4]thiazin-3-[(S)-2-ylamino]methyl]-4H-pyrido[3,2-b][1,4]thiazin-3-[(S)-2-ylamino]methyl]-4H-pyrido[3,2-b][1,4]thiazin-3-[(S)-2-ylamino]methyl]-4H-pyrido[3,2-b][1,4]thiazin-3-[(S)-2-ylamino]methyl]-4H-pyrido[3,2-b][1,4]thiazin-3-[(S)-2-ylamino]methyl]-4H-pyrido[3,2-b][1,4]thiazin-3-[(S)-2-ylamino]methyl]-4H-pyrido[3,2-b][1,4
- hydroxy-2-(6-methoxyquinolin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4*H*-pyrido[3,2-35] b][1,4]thiazin-3-one;
- 6-({(3R,4S)-1-[2-(2,3-Dihydro-[1,4]dioxino[2,3-f]quinolin-10-yl)-ethyl]-3-fluoro-piperidin-4-ylamino}-methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one and 6-({(3S,4R)-1-[2-

(2,3-dihydro-[1,4]dioxino[2,3-f]quinolin-10-yl)-ethyl]-3-fluoro-piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one;
6-({(3R,4S)-1-[2-(6,8-Difluoro-quinolin-4-yl)-ethyl]-3-fluoro-piperidin-4-ylamino}methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one and 6-({(3S,4R)-1-[2-(6,8-difluoro-quinolin4-yl)-ethyl]-3-fluoro-piperidin-4-ylamino}-methyl)-4H-pyrido[3,2-b][1,4]thiazin-3-one;
6-[({(3S,4R)-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxy-[1,5]naphthyridin-4-yl)ethyl]piperidin-4-ylamino}methyl)-4H-benzo[1,4]thiazin-3-one and 6-[({(3R,4S)-3Fluoro-1-[(R)-2-hydroxy-2-(6-methoxy-[1,5]naphthyridin-4-yl)-ethyl]piperidin-4ylamino}methyl)-4H-benzo[1,4]thiazin-3-one;

- 6-[({cis-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxy-[1,5]naphthyridin-4-yl)-ethyl]piperidin-4-ylamino}methyl)-4H-benzo[1,4]thiazin-3-one Faster running Diastereoisomer;
 - 6-[($\{(cis-3-Fluoro-1-[(R)-2-hydroxy-2-(6-methoxy-[1,5]naphthyridin-4-yl)-ethyl]$ piperidin-4-ylamino $\}$ methyl)-4H-benzo[1,4]thiazin-3-one Slower-running
- Diastereoisomer;

6- $(\{2S,4S\}-1-[(R)-2-hydroxy-2-(6-methoxy-[1,5]naphthyridin-4-yl)ethyl]-2-(trifluoromethyl)piperidin-4-ylamino\} methyl)-4H-pyrido[1,4]thiazin-3-one; 6-<math>(\{2S,4R\}-1-[(R)-2-hydroxy-2-(6-methoxy-[1,5]naphthyridin-4-yl)ethyl]-2-(trifluoromethyl)piperidin-4-ylamino\} methyl)-4H-pyrido[1,4]thiazin-3-one;$

- 20 or a pharmaceutically acceptable derivative thereof.
 - 11. A method of treatment of bacterial infections in mammals, particularly in man, which method comprises the administration to a mammal in need of such treatment an effective amount of a compound according to claim 1.

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- 12. The use of a compound according to claim 1, in the manufacture of a medicament for use in the treatment of bacterial infections in mammals.
- 13. A pharmaceutical composition comprising a compound according to claim 1 and a pharmaceutically acceptable carrier for use in the treatment of bacterial infections in mammals.
 - 14. A pharmaceutical composition comprising a compound according to claim 1, and a pharmaceutically acceptable carrier.

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15. A compound according to claim 1 for use as a medicament.

- 16. A compound according to claim 1 for use in the treatment of bacterial infections in mammals.
- 17. A process for preparing a compound of formula (I) according to claim 1, or a pharmaceutically acceptable derivative thereof, which process comprises reacting a compound of formula (IV) with a compound of formula (V):

$$R^{1'}$$
 $Z^{2'}$
 $Z^{3'}$
 $Z^{4'}$
 $Z^{4'}$
 $Z^{5'}$
 $Z^{4'}$
 $Z^{5'}$
 $Z^{4'}$
 $Z^{5'}$
 $Z^{5'}$

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wherein n is as defined in formula (I); $Z^{1'}$, $Z^{2'}$, $Z^{3'}$, $Z^{4'}$, $Z^{5'}$, $R^{1'}$, and $R^{3'}$ are Z^{1} , Z^{2} , Z^{3} , Z^{4} , Z^{5} , R^{1} , and R^{3} as defined in formula (I) or groups convertible thereto; Q^{1} is $NR^{2'}R^{4'}$ or a group convertible thereto wherein $R^{2'}$ and $R^{4'}$ are R^{2} and R^{4} as defined in formula (I) or groups convertible thereto and Q^{2} is H or $R^{3'}$ or Q^{1} and Q^{2} together form an optionally protected oxo group;

- (i) X is A'-COW, Y is H and n is 0;
- (ii) X is $CR^6=CR^8R^9$, Y is H and n is 0;
- (iii) X is oxirane, Y is H and n is 0;
- (iv) X is N=C=O and Y is H and n is 0;
- 20 (v) one of X and Y is CO₂Ry and the other is CH₂CO₂R^x;
 - (vi) X is CHR^6R^7 and Y is $C(=0)R^9$;
 - (vii) X is $CR^7=PR^2_3$ and Y is $C(=0)R^9$ and n=1;
 - (viii) X is $C(=0)R^7$ and Y is $CR^9=PR^2_3$ and n=1;
 - (ix) Y is COW and X is NHR 11 ' or NR11'COW and n=0 or 1 or when n=1 X is COW and Y is NHR 11 ' or NR11'COW:
 - (x) X is NHR^{11'} and Y is $C(=0)R^8$ and n=1;
 - (xi) X is NHR^{11'} and Y is CR^8R^9W and n=1;
 - (xii) X is NR¹¹'COCH₂W or NR¹¹'SO₂CH₂W and Y is H and n=0;
 - (xiii) $X \text{ is } CR^6R^7SO_2W \text{ and } Y \text{ is } H \text{ and } n=0;$
- 30 (xiv) X is W or OH and Y is CH₂OH and n is 1;
 - (xv) X is NHR^{11} ' and Y is SO_2W or X is NR^{11} ' SO_2W and Y is H, and n is 0;
 - (xvi) X is W and Y is CONHR¹¹;

in which W is a leaving group, e.g. halo or imidazolyl; R^x and R^y are (C_{1-6}) alkyl; R^z is aryl or (C_{1-6}) alkyl; A' and NR^{11} ' are A and NR^{11} as defined in formula (I), or groups convertible thereto; and oxirane is:

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wherein R^6 , R^8 and R^9 are as defined in formula (I); and thereafter optionally or as necessary converting Q^1 and Q^2 to NR^2R^4 ; converting A', Z^1 ', Z^2 ', Z^3 ', Z^4 ', Z^5 ', R^1 ', R^2 ', R^3 ', R^4 and NR^{11} '; to A, Z^1 , Z^2 , Z^3 , Z^4 , Z^5 , R^1 , R^2 , R^3 , R^4 and NR^{11} ; converting A-B to other A-B, interconverting R^1 , R^2 , R^3 and/or R^4 , and/or forming a pharmaceutically acceptable derivative thereof.